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# FACULTY OF MEDICAL SCIENCES

## UNIT-I Topic:- Cell-mediated immunity.



Involves specialized set of lymphocytes called T cells that recognize foreign antigens on the surface of cells, organisms, or tissues:

Two types of T cells are present:Helper T cellsCytotoxic T cells

 T cells regulate proliferation and activity of other cells of the immune system: B cells, macrophages, neutrophils, etc.

 Defense against: Bacteria and viruses that are inside host cells and are inaccessible to antibodies and also to Fungi, protozoa, and helminthes also to Cancer cells and Transplanted tissues.

#### **CELL MEDIATED IMMUNE RESPONSES**

Primary Function Of Cell Mediated Response:-

- •Eliminate Intracellular Pathogens
- •Eliminate Tumor Cells
- Both Ag Specific And Non-specific cells Are Involved
- •Ag Specific: CD8+ Cells (TC) And TH (DTH)
- •Non-specific: Neutrophils, NK
- Both Specific And Non-specific Require Cytokines
- •Humoral And Cell Mediated Do Collaborate

#### CMI may play a role in some harmful conditions:

- Hypersensitivity reactions type IV (contact dermatitis)
- Graft rejection
- Autoimmune diseases
- Cell mediated cytotoxicity mediated by:
- T-Cytotoxic cells.
- Natural killer cells.
- Activated macrophages.



### **CMI HELPS IN**

Delayed hypersensitivity

•Immunity in infections caused by Obligate and facultative intracellular parasites

•E.g. – Tuberculosis, Leprosy, Listeriosis, Brucellosis,
•Fungi – Histoplasmosis, Cocccidiomysosis, Blastomycosis,
•Parasites – Trypanosomiasis

•In transplantation immunity,



- •Pathogenesis of Autoimmune diseases
- •Cell Mediated Immunity Can Be Divided Into 2 Major Categories
- •Effectors lyse target
- •2 groups of cells: CTLs (specific) and NK.
- •Effectors which are CD4+ and mediate DTH

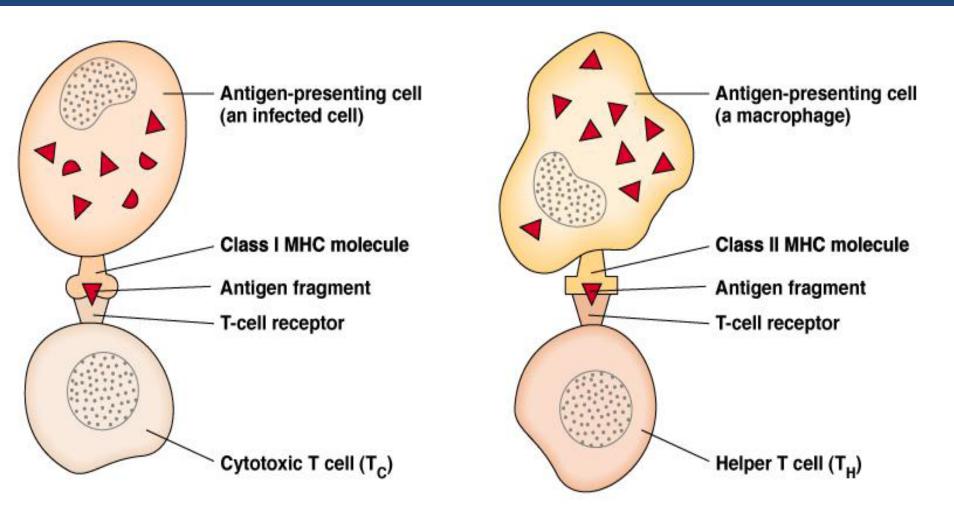
Cellular Components of Immunity:

- •T cells are key cellular component of immunity.
- •T cells have an antigen receptor that recognizes and reacts to a specific antigen (T cell receptor).
- •T cell receptor only recognize antigens combined with major histocompatibility (MHC) proteins on the surface of cells.

MHC Class I: Found on all cells.
 MHC Class II: Found on phagocytes.

•Clonal selection increases number of T cells.

#### T CELLS ONLY RECOGNIZE ANTIGEN ASSOCIATED WITH MHC MOLECULES ON CELL SURFACES



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### HOW CTLS KILL

### Phases In CTL Killing

- •Conjugate formation.
- •LFA-1 (CTL) binds ICAMs (Target).
- •LFA-1 changes to high avidity if Ag Is Recognized.
- •Activated LFA-1 persists for 5-10 mins.
- •Membrane attack.
- •Requires Ca2+ and energy
- •Granules release Perforins (65 kDa) and Granzymes (serine proteases) at the junctional space.
- •Perforins polymerize forming cylindrical pores (5-20 nm), Ca2+ is needed.
- •Granzymes enter target cell.
- •Granzyme B can enter thru mannose-6-phosphate receptor in a vesicle.
- •DNA fragmentation.
- •CTL dissociation.
- •Target cell destruction.
- •Apoptotic death within a few hours.